

10/533081

Connecting via Winsock to STN

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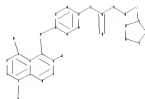
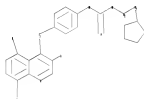
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FILE 'HOME' ENTERED AT 14:19:04 ON 01 JUL 2009

=> file reg

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Uploading C:\Program Files\Stnexp\Queries\10533081.str



```
chain nodes :
22 24 25 26 27 28 29 30 31 32
ring nodes :
1 2 3 4 5 6 7 8 9 10 11 12 13 14 15 16 17 18 19 20 21
chain bonds :
2-25 5-24 7-22 8-26 12-22 15-27 21-31 27-28 28-29 28-32 29-30 30-31
ring bonds :
1-2 1-6 1-10 2-3 3-4 4-5 5-6 6-7 7-8 8-9 9-10 11-12 11-16 12-13 13-14
14-15 15-16 17-18 17-21 18-19 19-20 20-21
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10/533081

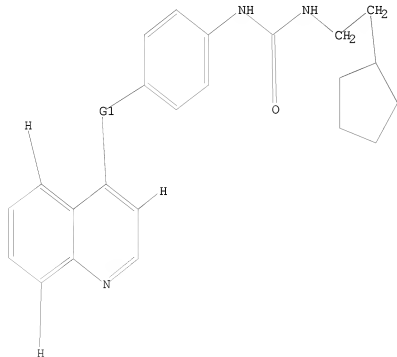
exact/norm bonds :
7-22 12-22 15-27 27-28 28-29 28-32
exact bonds :
2-25 5-24 8-26 17-18 17-21 18-19 19-20 20-21 21-31 29-30 30-31
normalized bonds :
1-2 1-6 1-10 2-3 3-4 4-5 5-6 6-7 7-8 8-9 9-10 11-12 11-16 12-13 13-14
14-15 15-16
isolated ring systems :
containing 1 : 11 : 17 :

G1:O,S

Match level :
1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:Atom 8:Atom 9:Atom 10:Atom
11:Atom 12:Atom 13:Atom 14:Atom 15:Atom 16:Atom 17:Atom 18:Atom 19:Atom
20:Atom 21:Atom 22:CLASS 24:CLASS 25:CLASS 26:CLASS 27:CLASS 28:CLASS
29:CLASS 30:CLASS 31:CLASS 32:CLASS

L1 STRUCTURE UPLOADED

=> d l1
L1 HAS NO ANSWERS
L1 STR



G1 O,S

Structure attributes must be viewed using STN Express query preparation.

=> s ll sam

SAMPLE SEARCH INITIATED 14:20:02 FILE 'REGISTRY'

SAMPLE SCREEN SEARCH COMPLETED - 19 TO ITERATE

100.0% PROCESSED 19 ITERATIONS

0 ANSWERS

SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **COMPLETE**

BATCH **COMPLETE**

PROJECTED ITERATIONS: 119 TO 641

PROJECTED ANSWERS: 0 TO 0

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=> s ll full

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100.0% PROCESSED 313 ITERATIONS

3 ANSWERS

SEARCH TIME: 00.00.01

L3 3 SEA SSS FUL L1

=> file caplus

=> s l3

L4 1 L3

=> d ibib abs hitstr

L4 ANSWER 1 OF 1 CAPLUS COPYRIGHT 2009 ACS on STN

ACCESSION NUMBER: 2004:390229 CAPLUS

DOCUMENT NUMBER: 140:406748

TITLE: Preparation of quinoline derivatives and quinazoline derivatives inhibiting autophosphorylation of Flt3 and medicinal compositions containing the same

INVENTOR(S): Hirai, Hisamaru; Miwa, Atsushi; Yoshino, Tetsuya; Kurokawa, Mineo

PATENT ASSIGNEE(S): Kirin Beer Kabushiki Kaisha, Japan; Hirai, Naoko

SOURCE: PCT Int. Appl., 118 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: Japanese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|-----------------|----------|
| WO 2004039782 | A1 | 20040513 | WO 2003-JP13848 | 20031029 |
| W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW | | | | |

RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG

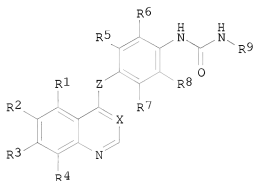
AU 2003280599 A1 20040525 AU 2003-280599 20031029
 EP 1566379 A1 20050824 EP 2003-769958 20031029

R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK

US 20080207617 A1 20080828 US 2006-533081 20061109
 JP 2002-314670 A 20021029
 WO 2003-JP13848 W 20031029

PRIORITY APPLN. INFO.:

OTHER SOURCE(S): MARPAT 140:406748
 GI



I

AB Disclosed is a medicinal composition to be used in preventing or treating diseases which can be effectively treated or prevented by inhibiting autophosphorylation of Flt3, comprising a compound represented by the following general formula (I) or pharmaceutically acceptable salts thereof or solvates of the same [wherein X = CH, N; Z = O, S; R1, R2, R3 = H, OH, halo, NO2, cyano, CHO, or each optionally substituted NH2, C1-6 alkyl, C2-6 alkenyl, C2-6 alkynyl, C1-6 alkoxy, C1-4 alkyl-carbonyl, or CONH2; R4 = H; R5 = R6 = R7 = R8 = H, or 1 or 2 number of R5-R8 = halo, C1-4 alkyl, C1-4 alkoxy, NO2, NH2, OH and all the others = H; R9 = (a) saturated 3- to 9-membered carbocyclyl optionally substituted by 1-3 number of C1-4 alkyl or (b) C1-4 alkyl substituted by C1-4 alkoxy, 5- or 6-membered heterocyclyl, each (un)substituted saturated 3- to 9-membered carbocyclyl, iso-Pr, tert-Bu, or NH2]. The diseases which can be effectively treated by inhibiting autophosphorylation of Flt3 include hematopoietic malignant tumor, in particular acute myelocytic leukemia or bone marrow neoplastic abnormality syndrome. Thus, 2 g 4-[(6,7-dimethoxy-4-quinolinyl)oxy]aniline was dissolved in 100 mL CHCl3, treated dropwise with a solution of 2 mL Et3N and 1 g triphosgene in 4 mL CHCl3, stirred at room temperature for 30 min, treated with 750 mg 3,3-dimethylbutylamine, and stirred at room temperature for 5 h to give, after workup and silica gel chromatog., N-[4-[(6,7-dimethoxy-4-quinolinyl)oxy]phenyl]-N'-(3,3-dimethylbutyl)urea (II). II.HCl and N-[4-[(6,7-dimethoxy-4-quinolinyl)oxy]-3-fluorophenyl]-N'-(3,3-dimethylbutyl)urea hydrochloride showed IC50 of 2 and <1 nM, resp., for inhibiting the autophosphorylation of MV4-11 human leukemia cell.

IT 688309-27-5P 688309-49-1P 688309-61-7P

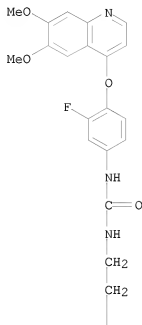
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of quinoline and quinazoline derivs. as inhibitors of autophosphorylation of FMS-like tyrosine kinase 3 (Flt3) for treatment or preparation of hematopoietic malignant tumor)

RN 688309-27-5 CAPLUS

CN Urea, N-(2-cyclopentylethyl)-N'-[4-[(6,7-dimethoxy-4-quinolinyl)oxy]-3-fluorophenyl]- (CA INDEX NAME)

PAGE 1-A



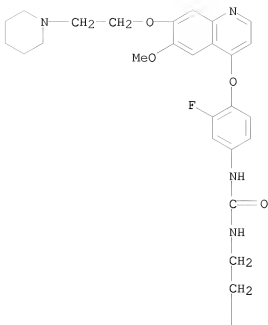
PAGE 2-A



RN 688309-49-1 CAPLUS

CN Urea, N-(2-cyclopentylethyl)-N'-[3-fluoro-4-[[6-methoxy-7-[2-(1-piperidinyl)ethoxy]-4-quinolinyl]oxy]phenyl]- (CA INDEX NAME)

PAGE 1-A



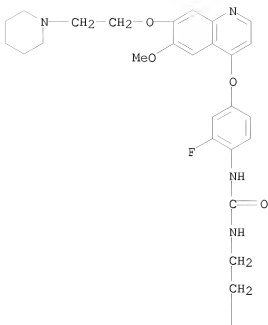
PAGE 2-A



RN 688309-61-7 CAPLUS

CN Urea, N-(2-cyclopentylethyl)-N'-[2-fluoro-4-[[6-methoxy-7-[2-(1-piperidinyl)ethoxy]-4-quinolinyloxy]phenyl]- (CA INDEX NAME)

PAGE 1-A



PAGE 2-A



REFERENCE COUNT: 9 THERE ARE 9 CITED REFERENCES AVAILABLE FOR THIS
RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

=> file marpat

=> s 13 full

FULL SEARCH INITIATED 14:20:25 FILE 'MARPAT'

FULL SCREEN SEARCH COMPLETED - 6395 TO ITERATE

100.0% PROCESSED 6395 ITERATIONS

2 ANSWERS

SEARCH TIME: 00.00.05

L5 2 SEA SSS FUL L1

=> d his

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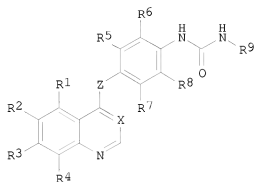
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L5 ANSWER 1 OF 2 MARPAT COPYRIGHT 2009 ACS on STN
 ACCESSION NUMBER: 140:406748 MARPAT
 TITLE: Preparation of quinoline derivatives and quinazoline
 derivatives inhibiting autophosphorylation of Flt3 and
 medicinal compositions containing the same
 INVENTOR(S): Hirai, Hisamaru; Miwa, Atsushi; Yoshino, Tetsuya;
 Kurokawa, Mineo
 PATENT ASSIGNEE(S): Kirin Beer Kabushiki Kaisha, Japan; Hirai, Naoko
 SOURCE: PCT Int. Appl., 118 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: Japanese
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------------------------|--|----------|-----------------|----------|
| WO 2004039782 | A1 | 20040513 | WO 2003-JP13848 | 20031029 |
| W: | AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW | | | |
| RW: | GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG | | | |
| AU 2003280599 | A1 | 20040525 | AU 2003-280599 | 20031029 |
| EP 1566379 | A1 | 20050824 | EP 2003-769958 | 20031029 |
| R: | AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK | | | |
| US 20080207617 | A1 | 20080828 | US 2006-533081 | 20061109 |
| PRIORITY APPLN. INFO.: | | | JP 2002-314670 | 20021029 |
| | | | WO 2003-JP13848 | 20031029 |

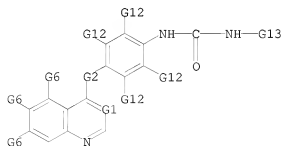
GI



I

AB Disclosed is a medicinal composition to be used in preventing or treating diseases which can be effectively treated or prevented by inhibiting autophosphorylation of Flt3, comprising a compound represented by the following general formula (I) or pharmaceutically acceptable salts thereof or solvates of the same [wherein X = CH, N; Z = O, S; R1, R2, R3 = H, OH, halo, NO2, cyano, CHO, or each optionally substituted NH2, C1-6 alkyl, C2-6 alkenyl, C2-6 alkynyl, C1-6 alkoxy, C1-4 alkyl-carbonyl, or CONH2; R4 = H; R5 = R6 = R7 = R8 = H, or 1 or 2 number of R5-R8 = halo, C1-4 alkyl, C1-4 alkoxy, NO2, NH2, OH and all the others = H; R9 = (a) saturated 3- to 9-membered carbocyclyl optionally substituted by 1-3 number of C1-4 alkyl or (b) C1-4 alkyl substituted by C1-4 alkoxy, 5- or 6-membered heterocyclyl, each (un)substituted saturated 3- to 9-membered carbocyclyl, iso-Pr, tert-Bu, or NH2]. The diseases which can be effectively treated by inhibiting autophosphorylation of Flt3 include hematopoietic malignant tumor, in particular acute myelocytic leukemia or bone marrow neoplastic abnormality syndrome. Thus, 2 g 4-[6,7-dimethoxy-4-quinolinyl]oxylaniline was dissolved in 100 mL CHCl3, treated dropwise with a solution of 2 mL Et3N and 1 g triphosgene in 4 mL CHCl3, stirred at room temperature for 30 min, treated with 750 mg 3,3-dimethylbutylamine, and stirred at room temperature for 5 h to give, after workup and silica gel chromatog., N-[4-[(6,7-dimethoxy-4-quinolinyl)oxy]phenyl]-N'-(3,3-dimethylbutyl)urea (II). II.HCl and N-[4-[(6,7-dimethoxy-4-quinolinyl)oxy]-3-fluorophenyl]-N'-(3,3-dimethylbutyl)urea hydrochloride showed IC50 of 2 and <1 nM, resp., for inhibiting the autophosphorylation of MV4-11 human leukemia cell.

MSTR 1



G1 = CH
G2 = O
G13 = 89



G32 = (2-4) CH2

Patent location:

Note: claim 1
or pharmaceutically acceptable salts or solvates
Note: additional ring formation also claimed

REFERENCE COUNT: 9 THERE ARE 9 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L5 ANSWER 2 OF 2 MARPAT COPYRIGHT 2009 ACS on STN

ACCESSION NUMBER: 140:357353 MARPAT

TITLE: Preparation of triazolone and triazoloethione inhibitors of matrix metalloproteinases and/or TNF- α converting enzyme as anti-inflammatory agents

INVENTOR(S): King, Bryan W.; Sheppeck, James; Gilmore, John L.

PATENT ASSIGNEE(S): Bristol-Myers Squibb Company, USA

SOURCE: PCT Int. Appl., 125 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

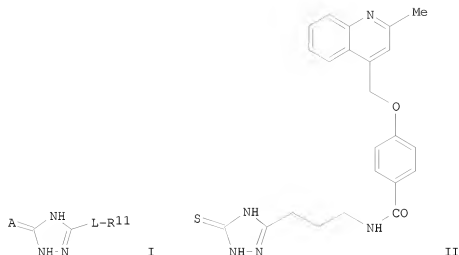
LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

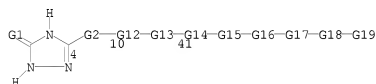
| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
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| WO 2004032846 | A2 | 20040422 | WO 2003-US31537 | 20031003 |
| WO 2004032846 | A3 | 20040715 | | |
| W: | AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW | | | |
| RW: | GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG | | | |
| AU 2003284001 | A1 | 20040504 | AU 2003-284001 | 20031003 |
| US 20040116491 | A1 | 20040617 | US 2003-678331 | 20031003 |
| US 7074810 | B2 | 20060711 | | |
| EP 1558581 | A2 | 20050803 | EP 2003-776228 | 20031003 |
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| PRIORITY APPLN. INFO.: | | | US 2002-416709P | 20021007 |
| | | | WO 2003-US31537 | 20031003 |

GI



AB The present application describes novel hydantoin derivs. (shown as I; A = O, S; L-R11 represents a very large variety of substituents and is defined in the claims; e.g. II) or pharmaceutically acceptable salt or prodrug forms thereof, which are useful as inhibitors of matrix metalloproteinases (MMP), TNF- α converting enzyme (TACE), aggrecanase, or a combination thereof. Some examples of I exhibit K_i 's $<10 \mu\text{M}$ but individual data are not presented. Although the methods of preparation are not claimed, 37 example preps. are included. For example, II was prepared in 4 steps (100, 66, 73 and 82%, resp.) starting with condensation of Et 4-aminobutyrate hydrochloride with 4-(2-methylquinolin-4-ylmethoxy)benzoyl chloride hydrochloride followed by base hydrolysis to the acid, followed by hydrazide formation with thiosemicarbazide followed by cyclization.

MSTR 1



G2 = 31-4 32-10



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G10 = 108



G12 = carbon chain <containing 1 or more C,
0 or more double bonds, 0 or more triple bonds>
(opt. substd.)

G13 = 89-10 90-41



G14 = bond

G15 = NH (opt. substd.)

G16 = phenylene (opt. substd.)

G17 = O

G18 = bond

G19 = quinoliny (opt. substd. by G23)

Patent location: claim 1

Note: or pharmaceutically acceptable salts

Note: additional ring formation also claimed

Note: substitution is restricted

Stereochemistry: or stereoisomers

REFERENCE COUNT: 1 THERE ARE 1 CITED REFERENCES AVAILABLE FOR THIS
RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

=> d his

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FILE 'REGISTRY' ENTERED AT 14:19:45 ON 01 JUL 2009

L1 STRUCTURE UPLOADED

L2 0 S L1 SAM

L3 3 S L1 FULL

FILE 'CAPLUS' ENTERED AT 14:20:10 ON 01 JUL 2009

L4 1 S L3

FILE 'MARPAT' ENTERED AT 14:20:19 ON 01 JUL 2009

L5 2 S L3 FULL

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Executing the logoff script...

10/533081

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